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NEWS 8 SEP 22 MATHDI to be removed from STN

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MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 13 JUNE 2005

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NEWS INTER General Internet Information

NEWS LOGIN Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN

NEWS WWW CAS World Wide Web Site (general information)

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=> file reg

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 0.21 0.21

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STRUCTURE FILE UPDATES: 25 SEP 2005 HIGHEST RN 863878-84-6 DICTIONARY FILE UPDATES: 25 SEP 2005 HIGHEST RN 863878-84-6

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Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=> Uploading C:\Program Files\Stnexp\Queries\10743287.str

chain nodes : 23 10 11 12 13 14 15 16 17 18 19 20 1 2 3 4 5 6 7 26 27 28 chain bonds : 1-2 1-16 1-18 2-3 2-25 2-26 3-4 4-5 5-6 6-7 7-8 8-9 8-20 9-10 10-11 11-12 12-13 13-14 14-15 14-27 14-28 15-17 15-19 18-23 19-22 exact/norm bonds : 1-16 1-18 8-20 15-17 15-19 18-23 19-22 exact bonds : 1-2 2-3 2-25 2-26 3-4 4-5 5-6 6-7 7-8 8-9 9-10 10-11 11-12 13-14 14-15 14-27 14-28

G1:H,Ak

Match level:
1:CLASS 2:CLASS 3:CLASS 4:CLASS 5:CLASS 6:CLASS 7:CLASS 8:CLASS 9:CLASS
10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS
18:CLASS 19:CLASS 20:CLASS 22:CLASS 23:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS
Page 2

L1 STRUCTURE UPLOADED

=> s l1

SAMPLE SEARCH INITIATED 17:07:29 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 581 TO ITERATE

100.0% PROCESSED 581 ITE

581 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

10174 TO 13066

PROJECTED ANSWERS:

0 TO

L2

0 SEA SSS SAM L1

=> s l1 ful

FULL SEARCH INITIATED 17:07:35 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 11481 TO ITERATE

100.0% PROCESSED 11481 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

T.3 2 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY 161.33 SESSION

161.54

FULL ESTIMATED COST

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FILE COVERS 1907 - 26 Sep 2005 VOL 143 ISS 14 FILE LAST UPDATED: 25 Sep 2005 (20050925/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 13

L4

2 L3

=> d l4 ibib hitstr abs 1-2

L4 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:648491 CAPLUS

DOCUMENT NUMBER: 141:190505

TITLE: Preparation of hydroxyl compounds for cholesterol

management and related uses

INVENTOR(S): Dasseux, Jean-Louis Henri; Oniciu, Daniela Carmen

PATENT ASSIGNEE(S): Esperion Therapeutics, Inc., USA

SOURCE: PCT Int. Appl., 348 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT 1	NO.		DATE	APPLICATION NO.	DATE
					2002122
				WO 2003-US41411	20031223
	067489				
WO 2004	067489				
W:	AE, AG, AL	, AM, AT,	, AU, AZ,	BA, BB, BG, BR, BY,	BZ, CA, CH, CN,
	CO, CR, CU	, CZ, DE,	, DK, DM,	DZ, EC, EE, EG, ES,	FI, GB, GD, GE,
	GH, GM, HR	, HU, ID,	, IL, IN,	IS, JP, KE, KG, KP,	KR, KZ, LC, LK,
	LR, LS, LT	, LU, LV,	, MA, MD,	MG, MK, MN, MW, MX,	MZ, NI, NO, NZ,
	OM, PG, PH	, PL, PT	, RO, RU,	SC, SD, SE, SG, SK,	SL, SY, TJ, TM,
	TN, TR, TT	, TZ, UA,	, UG, US,	UZ, VC, VN, YU, ZA,	ZM, ZW
RW:	BW, GH, GM	, KE, LS,	, MW, MZ,	SD, SL, SZ, TZ, UG,	ZM, ZW, AM, AZ,
•	BY, KG, KZ	, MD, RU	, TJ, TM,	AT, BE, BG, CH, CY,	CZ, DE, DK, EE,
	ES, FI, FR	, GB, GR	, HU, IE,	IT, LU, MC, NL, PT,	RO, SE, SI, SK,
	TR. BF. BJ	, CF, CG	, CI, CM,	GA, GN, GQ, GW, ML,	MR, NE, SN, TD, TG
US 2004	209847	A1	20041021	US 2003-743287 US 2003-743109	20031223
US 2004	214887	A1	20041028	US 2003-743109 ≺	20031223
US 2005	043278	A1	20050224	US 2003-743470	20031223
PRIORITY APP				US 2003-441795∳	P 20030123
OTHER SOURCE	(S):				
IT 738606-	46-7P 8-HV	droxy-2.2	2.14.14-t	etramethylpentadecan	edioic acid
RI. PAC	(Pharmacol	ogical ac	ctivity):	SPN (Synthetic prepared)	aration); THU
(Theran	eutic use):	BTOL (B	iological	study); PREP (Prepa	ration); USES
(Uses)	Cucio ubo, ,				
(obcb)	icholestere	mic agent	t: prepar	ation of hydroxyalka	nes for cholesterol
	gement and				•
	46-7 CAPLU		,		
177 120000-	TO , CHELO		_		(OGT) (GN TNDEN

Pentadecanedioic acid, 8-hydroxy-2,2,14,14-tetramethyl- (9CI) (CA INDEX

TT 738606-64-9, 8-Hydroxy-2,2,14,14-tetramethylpentadecanedioic acid diethyl ester

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of hydroxyalkanes for cholesterol management and related uses)

CN

NAME)

CN Pentadecanedioic acid, 8-hydroxy-2,2,14,14-tetramethyl-, diethyl ester (9CI) (CA INDEX NAME)

GI

$$Y^{1}-(CH_{2})_{m}$$
 R^{1}
 X
 $(CH_{2})_{n}$
 R^{11}
 $(CH_{2})_{m}-Y^{2}$
 R^{12}
 R^{11}
 R^{1

Title hydroxyalkanes I [wherein m = 0-5; n = 3-7; X = (CH2)p or CH2; p = 0AB 0-4; R1, R2, R11, R12 = independently H, alkyl, alkenyl, alkynyl, Ph, PhCH2, wherein R1, R2, R11, and R12 are not simultaneously H; Y1, Y2 = independently alkyl, OH, CO2H, CO2R3, SO3H, (un) substituted heterocyclyl, (di)phosphate and triphosphate esters, etc.; R3 = (un)substituted alkyl, alkenyl, alkynyl, Ph, PhCH2; and pharmaceutically acceptable salts, hydrates, solvates, or mixts. thereof] were prepared for cholesterol management. Thus, reduction of 7-oxo-2,2,12,12-tetramethyltridecanedioic acid di-Et ester with Na(BH4) in MeOH gave 7-hydroxy-2,2,12,12tetramethyltridecanedioic acid di-Et ester (92%), which was saponified with KOH in EtOH to afford the diacid II (95%). The latter inhibited lipid synthesis in primary rat hepatocytes with IC50 of 3.4 μM . In addition, the present invention relates to I, compns. comprising I, and methods useful for treating and preventing a variety of diseases and conditions such as, but not limited to aging, Alzheimer's disease, cancer, cardiovascular disease, diabetic nephropathy, diabetic retinopathy, a disorder of glucose metabolism, dyslipidemia, dyslipoproteinemia, hypertension, impotence, inflammation, insulin resistance, lipid elimination in bile, obesity, oxysterol elimination in bile, pancreatitis, Parkinson's disease, a peroxisome proliferator activated receptor-associated disorder, phospholipid elimination in bile, renal disease, septicemia, metabolic syndrome disorders (e.g., Syndrome X), thrombotic disorder. Compds. and methods of the invention can also be used to modulate C reactive protein or enhance bile production in a patient (no data). certain embodiments, I, compns. comprising I, and methods of the invention are useful in combination therapy with other therapeutics, such as hypocholesterolemic and hypoglycemic agents.

L4 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:518921 CAPLUS

DOCUMENT NUMBER: 141:236191

TITLE: Effects of a novel dual lipid synthesis inhibitor and

its potential utility in treating dyslipidemia and

metabolic syndrome

AUTHOR(S): Cramer, Clay T.; Goetz, Brian; Hopson, Krista L. M.;

Fici, Gregory J.; Ackermann, Rose M.; Brown, Stephen C.; Bisgaier, Charles L.; Rajeswaran, W. G.; Oniciu,

Daniela C.; Pape, Michael E.

CORPORATE SOURCE:

Esperion Therapeutics, Inc., Ann Arbor, MI, 48108, USA

Journal of Lipid Research (2004), 45(7), 1289-1301

CODEN: JLPRAW; ISSN: 0022-2275

PUBLISHER: American Society for Biochemistry and Molecular

Biology, Inc.

DOCUMENT TYPE:

Journal English

LANGUAGE:

SOURCE:

IT

738606-46-7, ESP 55016

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(effects of lipid synthesis inhibitor and its potential utility in treating dyslipidemia and metabolic syndrome)

RN 738606-46-7 CAPLUS

CN Pentadecanedioic acid, 8-hydroxy-2,2,14,14-tetramethyl- (9CI) (CA INDEX NAME)

We have identified a novel ω -hydroxy-alkane dicarboxylic acid, ESP AB 55016, that favorably alters serum lipid variables in obese female Zucker (fa/fa) rats. ESP 55016 reduced serum non-HDL-cholesterol (non-HDL-C), triglyceride, and nonesterified fatty acid levels while increasing serum $\mathtt{HDL-C}$ and β -hydroxybutyrate levels in a dose-dependent manner. ESP 55016 reduced fasting serum insulin and glucose levels while also suppressing weight gain. In primary rat hepatocytes, ESP 55016 increased the oxidation of [14C] palmitate in a dose- and carnitine palmitoyl transferase-I (CPT-I) -dependent manner. Furthermore, in primary rat hepatocytes and in vivo, ESP 55016 inhibited fatty acid and sterol synthesis. The "dual inhibitor" activity of ESP 55016 was unlikely attributable to the activation of the AMP-activated protein kinase (AMPK) pathway because AMPK and acetyl-CoA carboxylase (ACC) phosphorylation states as well as ACC activity were not altered by ESP 55016. Further studies indicated the conversion of ESP 55016 to a CoA derivative in vivo. ESP 55016-CoA markedly inhibited the activity of partially purified ACC. The activity of partially purified HMG-CoA reductase was not altered by the xenobiotic-CoA. These data suggest that ESP 55016-CoA favorably alters lipid metabolism in a model of diabetic dyslipidemia in part by initially inhibiting fatty acid and sterol synthesis plus enhancing the oxidation of fatty acids through the ACC/malonyl-CoA/CPT-I regulatory axis.

REFERENCE COUNT: 70 THERE ARE 70 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> fil reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 11.23 172.77

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STRUCTURE FILE UPDATES: 25 SEP 2005 HIGHEST RN 863878-84-6 DICTIONARY FILE UPDATES: 25 SEP 2005 HIGHEST RN 863878-84-6

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TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2005

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Structure search iteration limits have been increased. See HELP SLIMITS for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

Uploading C:\Program Files\Stnexp\Queries\107432871.str

chain nodes:
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 22 23
chain bonds:
1-2 1-16 1-18 2-3 3-4 4-5 5-6 6-7 7-8 8-9 8-20 9-10 10-11 11-12 12-13
13-14 14-15 15-17 15-19 18-23 19-22
exact/norm bonds:
1-16 1-18 8-20 15-17 15-19 18-23 19-22
exact bonds:

1-2 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13 13-14 14-15

G1:H,Ak

Match level :

1:CLASS 2:CLASS 3:CLASS 4:CLASS 5:CLASS 6:CLASS 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 22:CLASS 23:CLASS

L5 STRUCTURE UPLOADED

=> s 15

SAMPLE SEARCH INITIATED 17:09:56 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 12887 TO ITERATE

15.5% PROCESSED 2000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED) SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 250940 TO 264540 PROJECTED ANSWERS: 0 TO 0

L6 0 SEA SSS SAM L5

=> s 15 ful

FULL SEARCH INITIATED 17:10:03 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 254675 TO ITERATE

98.2% PROCESSED 250033 ITERATIONS

5 ANSWERS

6 ANSWERS

O ANSWERS

100.0% PROCESSED 254675 ITERATIONS

SEARCH TIME: 00.00.18

6 SEA SSS FUL L5

=> d 17 ibib hitstr abs 1-6

'IBIB' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'
'HITSTR' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'
'ABS' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'

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REG - RN

SAM - Index Name, MF, and structure - no RN
FIDE - All substance data, except sequence data

IDE - FIDE, but only 50 names SOIDE - IDE, plus sequence data

SQIDE3 - Same as SQIDE, but 3-letter amino acid codes are used

SOD - Protein sequence data, includes RN

SOD3 - Same as SQD, but 3-letter amino acid codes are used

SQN - Protein sequence name information, includes RN

CALC - Table of calculated properties EPROP - Table of experimental properties

PROP - EPROP and CALC

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ABS -- Abstract

APPS -- Application and Priority Information

BIB -- CA Accession Number, plus Bibliographic Data

CAN -- CA Accession Number

CBIB -- CA Accession Number, plus Bibliographic Data (compressed)

IND -- Index Data

IPC -- International Patent Classification

PATS -- PI, SO

STD -- BIB, IPC, and NCL

IABS -- ABS, indented, with text labels IBIB -- BIB, indented, with text labels

ISTD -- STD format, indented

OBIB ----- AN, plus Bibliographic Data (original)

OIBIB ----- OBIB, indented with text labels

SBIB ----- BIB, no citations SIBIB ----- IBIB, no citations

The ALL format gives FIDE BIB ABS IND RE, plus sequence data when it is available.

The MAX format is the same as ALL.

The IALL format is the same as ALL with BIB ABS and IND indented, with text labels.

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=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 161.76 334.53

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL
ENTRY SESSION
CA SUBSCRIBER PRICE
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=> d his

L1

L5

(FILE 'HOME' ENTERED AT 17:07:03 ON 26 SEP 2005)

FILE 'REGISTRY' ENTERED AT 17:07:11 ON 26 SEP 2005

STRUCTURE UPLOADED

L2 0 S L1

L3 2 S L1 FUL

FILE 'CAPLUS' ENTERED AT 17:07:41 ON 26 SEP 2005

L4 2 S L3

FILE 'REGISTRY' ENTERED AT 17:09:40 ON 26 SEP 2005

STRUCTURE UPLOADED

L6 0 S L5

L7 6 S L5 FUL

FILE 'CAPLUS' ENTERED AT 17:10:59 ON 26 SEP 2005

=> s 17

L8 5 L7

=> d 18 ibib hitstr abs 1-5

L8 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2005:411319 CAPLUS

DOCUMENT NUMBER:

143:97083

TITLE:

Organozinc alkoxide-promoted aldol-Tishchenko reaction of aliphatic aldehydes: an expedient entry to prepare

the α -methylene ketones

AUTHOR(S):

Hon, Yung-Son; Chang, Chun-Ping

CORPORATE SOURCE:

Department of Chemistry and Biochemistry, National Chung Cheng University, Taipei, 621, Peop. Rep. China

SOURCE: Tetrahedron (2005), 61(22), 5267-5275

CODEN: TETRAB; ISSN: 0040-4020

PUBLISHER:

Elsevier B.V.

DOCUMENT TYPE:

Journal

LANGUAGE:

English

IT 856895-80-2P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of α -methylene ketones via organozinc alkoxide-promoted aldol-Tishchenko reaction of aliphatic aldehydes and subsequent oxidation

and

elimination)

RN 856895-80-2 CAPLUS

CN Pentadecanedioic acid, 8-hydroxy-7-[[(8-methoxy-1,8-dioxooctyl)oxy]methyl]-, dimethyl ester (9CI) (CA INDEX NAME)

AB I-ProZnEt is an excellent reagent to promote the aldol-Tishchenko reaction of the aliphatic aldehydes tethered with other labile functional groups. The 1,3-diol monoesters were formed as the major products, which could be converted to α-methylene ketones in two steps in good yields. E.g., reaction of BnCH2CHO in presence of i-ProZnEt gave BnCH2CH(OH)CHBnCH2O2CCH2Bn as the major product and BnCH2CH(O2CCH2Bn)CHBnCH2OH as the minor product. Oxidation of the mixture of products by PCC gave BnCH2COCHCH2O2CCH2Bn as the major product. Treatment of this ketone by DBU led to the α-methylene ketone BnCH2COCBn:CH2.

REFERENCE COUNT: 46 THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

2004:648491 CAPLUS

DOCUMENT NUMBER:

141:190505

TITLE:

Preparation of hydroxyl compounds for cholesterol

management and related uses

INVENTOR(S):

Dasseux, Jean-Louis Henri; Oniciu, Daniela Carmen

Esperion Therapeutics, Inc., USA

SOURCE:

PCT Int. Appl., 348 pp.

•

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT ASSIGNEE(S):

PA	TENT	NO.			KINI) 1	DATE		i	APPL:	ICAT:	ION 1	10.		DA	ATE		
	2004 2004				A2 A3		2004 2004		Ī	WO 2	003-1	US414	11		20	0312	223	
WO	2004				C1		2005											
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,	
		co.	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	GE,	
		GH.	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	ΚZ,	LC,	LK,	
		LR.	LS.	LT.	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NI,	NO,	NZ,	
		OM.	PG.	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	ТJ,	TM,	
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		BY.	KG.	KZ.	MD.	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	•
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·		TR,	BF,	ΒJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG
US	2004				Αĺ		2004	1021	•	US 2	003-	7432	87		2	0031	223	
US	2004	2148	87		A1		2004	1028		US 2	003-	7431	09		2	0031	223	
US	2005	0432	78		A1		2005	0224	•	US 2	003-	7434	70		2	0031	223	
PRIORIT	Y APP	LN.	INFO	.:		•			•	US 2	003-	4417	95P		P 2	0030	123	

OTHER SOURCE(S): MARPAT 141:190505

738606-46-7P, 8-Hydroxy-2,2,14,14-tetramethylpentadecanedioic acid 738606-61-6P, 8-Hydroxy-2,2,12,12-tetramethylpentadecanedioic acid diethyl ester

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(anticholesteremic agent; preparation of hydroxyalkanes for cholesterol management and related uses)

RN 738606-46-7 CAPLUS

CN Pentadecanedioic acid, 8-hydroxy-2,2,14,14-tetramethyl- (9CI) (CA_INDEX_NAME)

RN 738606-61-6 CAPLUS

CN Pentadecanedioic acid, 8-hydroxy-2,2,12,12-tetramethyl-, diethyl ester (9CI) (CA INDEX NAME)

738606-64-9, 8-Hydroxy-2,2,14,14-tetramethylpentadecanedioic acid diethyl ester

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of hydroxyalkanes for cholesterol management and related uses)

RN 738606-64-9 CAPLUS

CN Pentadecanedioic acid, 8-hydroxy-2,2,14,14-tetramethyl-, diethyl ester (9CI) (CA INDEX NAME)

O Me OH Me O
$$| | | | |$$
 EtO-C-C-(CH₂)₅-CH-(CH₂)₅-C-C-OEt Me Me

GI

Title hydroxyalkanes I [wherein m = 0-5; n = 3-7; X = (CH2)p or CH2; p = 0.5AB 0-4; R1, R2, R11, R12 = independently H, alkyl, alkenyl, alkynyl, Ph, PhCH2, wherein R1, R2, R11, and R12 are not simultaneously H; Y1, Y2 = independently alkyl, OH, CO2H, CO2R3, SO3H, (un) substituted heterocyclyl, (di)phosphate and triphosphate esters, etc.; R3 = (un)substituted alkyl, alkenyl, alkynyl, Ph, PhCH2; and pharmaceutically acceptable salts, hydrates, solvates, or mixts. thereof] were prepared for cholesterol management. Thus, reduction of 7-oxo-2,2,12,12-tetramethyltridecanedioic acid di-Et ester with Na(BH4) in MeOH gave 7-hydroxy-2,2,12,12tetramethyltridecanedioic acid di-Et ester (92%), which was saponified with KOH in EtOH to afford the diacid II (95%). The latter inhibited lipid synthesis in primary rat hepatocytes with IC50 of 3.4 μM. In addition, the present invention relates to I, compns. comprising I, and methods useful for treating and preventing a variety of diseases and conditions such as, but not limited to aging, Alzheimer's disease, cancer, cardiovascular disease, diabetic nephropathy, diabetic retinopathy, a disorder of glucose metabolism, dyslipidemia, dyslipoproteinemia, hypertension, impotence, inflammation, insulin resistance, lipid elimination in bile, obesity, oxysterol elimination in bile, pancreatitis, Parkinson's disease, a peroxisome proliferator activated receptor-associated disorder, phospholipid elimination in bile, renal disease, septicemia, metabolic syndrome disorders (e.g., Syndrome X), thrombotic disorder. Compds. and methods of the invention can also be used to modulate C reactive protein or enhance bile production in a patient (no data). certain embodiments, I, compns. comprising I, and methods of the invention are useful in combination therapy with other therapeutics, such as hypocholesterolemic and hypoglycemic agents.

ANSWER 3 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

2004:518921 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 141:236191

Effects of a novel dual lipid synthesis inhibitor and TITLE:

its potential utility in treating dyslipidemia and

metabolic syndrome

Cramer, Clay T.; Goetz, Brian; Hopson, Krista L. M.; AUTHOR (S):

Fici, Gregory J.; Ackermann, Rose M.; Brown, Stephen C.; Bisgaier, Charles L.; Rajeswaran, W. G.; Oniciu,

Daniela C.; Pape, Michael E.

CORPORATE SOURCE:

Esperion Therapeutics, Inc., Ann Arbor, MI, 48108, USA Journal of Lipid Research (2004), 45(7), 1289-1301 SOURCE:

CODEN: JLPRAW; ISSN: 0022-2275

American Society for Biochemistry and Molecular PUBLISHER:

Biology, Inc.

Journal DOCUMENT TYPE:

English LANGUAGE:

738606-46-7, ESP 55016 IT

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(effects of lipid synthesis inhibitor and its potential utility in treating dyslipidemia and metabolic syndrome)

RN 738606-46-7 CAPLUS

Pentadecanedioic acid, 8-hydroxy-2,2,14,14-tetramethyl- (9CI) (CA INDEX CN

We have identified a novel ω -hydroxy-alkane dicarboxylic acid, ESP AB 55016, that favorably alters serum lipid variables in obese female Zucker (fa/fa) rats. ESP 55016 reduced serum non-HDL-cholesterol (non-HDL-C), triglyceride, and nonesterified fatty acid levels while increasing serum HDL-C and β -hydroxybutyrate levels in a dose-dependent manner. ESP 55016 reduced fasting serum insulin and glucose levels while also suppressing weight gain. In primary rat hepatocytes, ESP 55016 increased the oxidation of [14C]palmitate in a dose- and carnitine palmitoyl transferase-I (CPT-I) -dependent manner. Furthermore, in primary rat hepatocytes and in vivo, ESP 55016 inhibited fatty acid and sterol synthesis. The "dual inhibitor" activity of ESP 55016 was unlikely attributable to the activation of the AMP-activated protein kinase (AMPK) pathway because AMPK and acetyl-CoA carboxylase (ACC) phosphorylation states as well as ACC activity were not altered by ESP 55016. Further studies indicated the conversion of ESP 55016 to a CoA derivative in vivo. ESP 55016-CoA markedly inhibited the activity of partially purified ACC. The activity of partially purified HMG-CoA reductase was not altered by the xenobiotic-CoA. These data suggest that ESP 55016-CoA favorably alters lipid metabolism in a model of diabetic dyslipidemia in part by initially inhibiting fatty acid and sterol synthesis plus enhancing the oxidation of fatty acids through the ACC/malonyl-CoA/CPT-I regulatory axis.

THERE ARE 70 CITED REFERENCES AVAILABLE FOR THIS 70 REFERENCE COUNT: RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 4 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

1987:196107 CAPLUS ACCESSION NUMBER:

106:196107

DOCUMENT NUMBER:

Enamine condensation on derivatives of aleuritic acid TITLE:

and synthesis of (Z)-9-tricosene (muscalure), its

(E) -isomer, and (E) -13-heptacosene

Subramanian, G. B. V.; Mehrotra, Alka; Mehrotra, AUTHOR (S):

Kalpana

Dep. Chem., Univ. Delhi, Delhi, 110007, India CORPORATE SOURCE:

Tetrahedron (1986), 42(14), 3967-72 SOURCE:

CODEN: TETRAB; ISSN: 0040-4020

Journal DOCUMENT TYPE: English LANGUAGE:

CASREACT 106:196107 OTHER SOURCE(S):

93416-11-6P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of, as intermediate in synthesis of heptacosene)

93416-11-6 CAPLUS RN

Pentadecanedioic acid, 7,8-dihydroxy-, (R*,R*)- (9CI) (CA INDEX NAME) CN

Relative stereochemistry.

$$_{\rm HO_2C}$$
 $_{\rm CH_2)_5}$ $_{\rm R}$ $_{\rm R}$ $_{\rm (CH_2)_6}$ $_{\rm CO_2H}$

Enamine condensation on (9RS, 10RS) -9, 10, 16-triacetoxyhexadecanoyl chloride AB as well as (7RS,8RS)-7,8-diacetoxy-1,15-pentadecadioyl chloride using 1-morpholino-1-cyclohexene led to chain elongated products with 22 and 27 carbon atoms resp. The 22 carbon product was converted into (Z)-9-tricosene and its E-isomer, while the 27 carbon product led to a synthesis of (E)-13-heptacosene.

ANSWER 5 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

1985:5642 CAPLUS

DOCUMENT NUMBER:

102:5642

TITLE:

Preparation of some hydroxypentadecane derivatives

from threo-aleuritic acid

AUTHOR (S):

SOURCE:

Subramanian, G. B. V.; Mehrotra, Kalpana

CORPORATE SOURCE:

Dep. Chem., Univ. Delhi, Delhi, 110 007, India Indian Journal of Chemistry, Section B: Organic

Chemistry Including Medicinal Chemistry (1984),

23B(4), 384-5

CODEN: IJSBDB; ISSN: 0376-4699

DOCUMENT TYPE:

Journal English

LANGUAGE:

93416-11-6P 93416-12-7P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of) 93416-11-6 CAPLUS

RN Pentadecanedioic acid, 7,8-dihydroxy-, (R*,R*)- (9CI) (CA INDEX NAME) CN

Relative stereochemistry.

93416-12-7 CAPLUS RN

Pentadecanedioic acid, 7,8-dihydroxy-, dimethyl ester, (R*,R*)- (9CI) INDEX NAME)

Relative stereochemistry.

MeO (CH₂)
$$\stackrel{\text{OH}}{\underset{\text{OH}}{\longrightarrow}}$$
 (CH₂) $\stackrel{\text{OMe}}{\underset{\text{OH}}{\longrightarrow}}$

The triacetate of threo-aleuritic acid (9,10,16-trihydroxypalmitic acid), on treatment with Pb(OAc)4 and iodine in the presence of light followed by deacetylation gave R(CH2)5CH(OH)CH(OH)(CH2)6R1 (I; R = CH2OH, R1 = CH2I) (II). Acetoxylation and hydrolysis of II gave I (R = R1 = CH2OH). II was reduced to give I (R = CH2OH, R1 = Me). Oxidation of protected I (R = CH2OH; R1 = CH2I, CH2OH, Me) gave the acids I (R = CO2H; R1 = CH2I, CO2H, Me), which were methylated to the resp. esters.

=> file req COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION 27.85 362.38 FULL ESTIMATED COST TOTAL SINCE FILE DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SESSION ENTRY -3.65 -5.11 CA SUBSCRIBER PRICE

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Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

Uploading C:\Program Files\Stnexp\Queries\107432872.str

chain nodes :

1 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 22 23

ring nodes : 2 25 26 chain bonds :

1-2 1-16 1-18 2-3 3-4 4-5 5-6 6-7 7-8 8-9 8-20 9-10 10-11 11-12 12-13 13-14 14-15 15-17 15-19 18-23 19-22

ring bonds :

2-25 2-26 25-26

exact/norm bonds :

1-16 1-18 2-25 2-26 8-20 15-17 15-19 18-23 19-22 25-26

exact bonds :

1-2 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13 13-14 14-15

G1:H, Ak

Match level : .

1:CLASS 2:CLASS 3:CLASS 4:CLASS 5:CLASS 6:CLASS 7:CLASS 8:CLASS 9:CLASS

10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 22:CLASS 23:CLASS 25:Atom 26:Atom

STRUCTURE UPLOADED L9

=> s 19

SAMPLE SEARCH INITIATED 17:15:34 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 449 TO ITERATE

100.0% PROCESSED

0 ANSWERS 449 ITERATIONS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

7709 TO 10251 PROJECTED ITERATIONS:

PROJECTED ANSWERS: 0 TO

0 SEA SSS SAM L9 L10

=> s 19 ful

FULL SEARCH INITIATED 17:15:39 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED -8839 TO ITERATE

100.0% PROCESSED 8839 ITERATIONS 1 ANSWERS

SEARCH TIME: 00.00.01

L11 1 SEA SSS FUL L9

=> file caplus

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
161.33
523.71

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION
CA SUBSCRIBER PRICE

0.00 -5.11

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=> s 111

L12 1 L11

=> d l12 ibib hitstr abs

L12 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2004:648491 CAPLUS

DOCUMENT NUMBER: 141:190505

TITLE: Preparation of hydroxyl compounds for cholesterol

management and related uses

INVENTOR(S): Dasseux, Jean-Louis Henri; Oniciu, Daniela Carmen

PATENT ASSIGNEE(S): Esperion Therapeutics, Inc., USA

SOURCE: PCT Int. Appl., 348 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004067489	A2	20040812	WO 2003-US41411	20031223
WO 2004067489	A3	20041125		
WO 2004067489	C1	20050217		
W: AE, AG, AL,	AM, AT	, AU, AZ, BA	A, BB, BG, BR, BY, BZ,	CA, CH, CN,

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CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE,
             GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK,
             LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,
             OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,
             TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
             TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                             US 2003-743287
     US 2004209847
                          A1
                                 20041021
                                                                     20031223
                                             US 2003-743109
                                                                     20031223
     US 2004214887
                          Α1
                                 20041028
                                             US 2003-743470
                                                                     20031223
     US 2005043278
                          A1
                                 20050224
                                             US 2003-441795P
                                                                     20030123
PRIORITY APPLN. INFO .:
OTHER SOURCE(S):
                         MARPAT 141:190505
    738607-05-1P, 13-(1-Carboxycyclopropyl)-8-hydroxy-2,2-
     dimethyltridecanoic acid
     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
        (anticholesteremic agent; preparation of hydroxyalkanes for cholesterol
        management and related uses)
RN
     738607-05-1 CAPLUS
     Cyclopropanetridecanoic acid, 1-carboxy-\eta-hydroxy-\alpha, \alpha-
CN
     dimethyl- (9CI) (CA INDEX NAME)
```

GΙ

Title hydroxyalkanes I [wherein m = 0-5; n = 3-7; X = (CH2)p or CH2; p = 0-4; R1, R2, R11, R12 = independently H, alkyl, alkenyl, alkynyl, Ph, PhCH2, wherein R1, R2, R11, and R12 are not simultaneously H; Y1, Y2 = independently alkyl, OH, CO2H, CO2R3, SO3H, (un)substituted heterocyclyl, (di)phosphate and triphosphate esters, etc.; R3 = (un)substituted alkyl, alkenyl, alkynyl, Ph, PhCH2; and pharmaceutically acceptable salts, hydrates, solvates, or mixts. thereof] were prepared for cholesterol management. Thus, reduction of 7-oxo-2,2,12,12-tetramethyltridecanedioic acid di-Et ester with Na(BH4) in MeOH gave 7-hydroxy-2,2,12,12-

tetramethyltridecanedioic acid di-Et ester (92%), which was saponified with KOH in EtOH to afford the diacid II (95%). The latter inhibited lipid synthesis in primary rat hepatocytes with IC50 of 3.4 µM. In addition, the present invention relates to I, compns. comprising I, and methods useful for treating and preventing a variety of diseases and conditions such as, but not limited to aging, Alzheimer's disease, cancer, cardiovascular disease, diabetic nephropathy, diabetic retinopathy, a disorder of glucose metabolism, dyslipidemia, dyslipoproteinemia, hypertension, impotence, inflammation, insulin resistance, lipid elimination in bile, obesity, oxysterol elimination in bile, pancreatitis, Parkinson's disease, a peroxisome proliferator activated receptor-associated disorder, phospholipid elimination in bile, renal disease, septicemia, metabolic syndrome disorders (e.g., Syndrome X), thrombotic disorder. Compds. and methods of the invention can also be used to modulate C reactive protein or enhance bile production in a patient (no data). certain embodiments, I, compns. comprising I, and methods of the invention are useful in combination therapy with other therapeutics, such as hypocholesterolemic and hypoglycemic agents.

=> file reg		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	6.29	530.00
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	-0.73	-5.84

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Uploading C:\Program Files\Stnexp\Queries\107432973.str

chain nodes :

1 3 4 5 6 7 8 9 10 11 12 13 15 16 17 18 19 20 22 23

ring nodes :

2 14 25 26 27 28

chain bonds :

1-2 1-16 1-18 2-3 3-4 4-5 5-6 6-7 7-8 8-9 8-20 9-10 10-11 11-12 12-13

13-14 14-15 15-17 15-19 18-23 19-22

ring bonds :

2-25 2-26 14-27 14-28 25-26 27-28

exact/norm bonds :

1-16 1-18 2-25 2-26 8-20 14-27 14-28 15-17 15-19 18-23 19-22 25-26

27-28

exact bonds :

1-2 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13 13-14 14-15

G1:H,Ak

Match level:

1:CLASS 2:CLASS 3:CLASS 4:CLASS 5:CLASS 6:CLASS 7:CLASS 8:CLASS 9:CLASS

10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 22:CLASS 23:CLASS 25:Atom 26:Atom 27:Atom

28:Atom

STRUCTURE UPLOADED L13

=> s 113

SAMPLE SEARCH INITIATED 17:18:01 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 230 TO ITERATE

100.0% PROCESSED 230 ITERATIONS O ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

3691 TO 5509 PROJECTED ITERATIONS:

O TO PROJECTED ANSWERS:

0 SEA SSS SAM L13 L14

=> s 113 ful

FULL SEARCH INITIATED 17:18:07 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 4730 TO ITERATE

100.0% PROCESSED 4730 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

L15 2 SEA SSS FUL L13

=> file caplus

SINCE FILE COST IN U.S. DOLLARS TOTAL SESSION ENTRY 161.33 691.33 FULL ESTIMATED COST TOTAL DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE SESSION ENTRY -5.84 CA SUBSCRIBER PRICE 0.00

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=> s 115

L16 1 L15

=> file reg

COST IN U.S. DOLLARS SINCE FILE TOTAL SESSION ENTRY FULL ESTIMATED COST 2.70 694.03 SINCE FILE TOTAL DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SESSION ENTRY 0.00 -5.84 CA SUBSCRIBER PRICE

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OH

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Uploading C:\Program Files\Stnexp\Queries\107432974.str

chain nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 17 18 19 20 21
chain bonds :
1-2 1-18 1-19 2-3 3-4 4-5 5-6 6-7 7-8 7-20 8-9 9-10 10-11 11-12 12-13
13-14 13-15 15-17 19-21
exact/norm bonds :
1-18 1-19 7-20 13-14 13-15 15-17 19-21
exact bonds :
1-2 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13

G1:H, Ak

Match level:
1:CLASS 2:CLASS 3:CLASS 4:CLASS 5:CLASS 6:CLASS 7:CLASS 8:CLASS 9:CLASS
10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 17:CLASS 18:CLASS
19:CLASS 20:CLASS 21:CLASS

STRUCTURE UPLOADED L17

=> s 117

SAMPLE SEARCH INITIATED 17:22:01 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 12887 TO ITERATE

2000 ITERATIONS 15.5% PROCESSED

INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.01

ONLINE **COMPLETE** FULL FILE PROJECTIONS:

> **COMPLETE** BATCH

264540 PROJECTED ITERATIONS: 250940 TO

0 TO PROJECTED ANSWERS:

0 SEA SSS SAM L17 L18

=> s l17 ful

FULL SEARCH INITIATED 17:22:09 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED -254675 TO ITERATE

98.1% PROCESSED 249891 ITERATIONS 5 ANSWERS 6 ANSWERS

0 ANSWERS

100.0% PROCESSED 254675 ITERATIONS

SEARCH TIME: 00.00.19

6 SEA SSS FUL L17 L19

=> file caplus

CA SUBSCRIBER PRICE

SINCE FILE TOTAL COST IN U.S. DOLLARS SESSION ENTRY 855.36 161.33 FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) TOTAL SINCE FILE SESSION ENTRY -5.84 0.00

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=> s 119

L20 3 L19

=> d l20 ibib hitstr abs 1-3

L20 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

2004:648491 CAPLUS

DOCUMENT NUMBER:

141:190505

TITLE:

Preparation of hydroxyl compounds for cholesterol

management and related uses

INVENTOR(S):

Dasseux, Jean-Louis Henri; Oniciu, Daniela Carmen

PATENT ASSIGNEE(S):

Esperion Therapeutics, Inc., USA

SOURCE:

PCT Int. Appl., 348 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE			
WO 2004067489 WO 2004067489 WO 2004067489	A3 20041125	WO 2003-US41411				
		BA, BB, BG, BR, BY,	RZ CA CH CN			
W: AE, AG, AL,	AM, AI, AU, AZ,	DZ, EC, EE, EG, ES,	FI GR GD GE			
CO, CR, CO,	CZ, DE, DR, DM,	TO TO VE VO VD	KD K7 I.C I.K			
GH, GM, HR,	HU, ID, IL, IN,	IS, JP, KE, KG, KP,	MZ NI NO NZ			
LR, LS, LT,	LU, LV, MA, MD,	MG, MK, MN, MW, MX,	CI CV TI TM			
OM, PG, PH,	PL, PT, RO, RO,	SC, SD, SE, SG, SK,	7M 7W			
TN, TR, TT,	TZ, UA, UG, US,	UZ, VC, VN, YU, ZA,	7M 7W AM A7			
RW: BW, GH, GM,	KE, LS, MW, MZ,	SD, SL, SZ, TZ, UG,	C7 DE DY EE			
BY, KG, KZ,	MD, RU, TJ, IM,	AT, BE, BG, CH, CY,	DO CE CI CK			
ES, FI, FR,	GB, GR, HU, IE,	IT, LU, MC, NL, PT,	MD NE CN TO TO			
TR, BF, BJ,	CF, CG, C1, CM,	GA, GN, GQ, GW, ML,	20031223			
US 2004209847	A1 20041021	US 2003-743287	20031223			
US 2004214887	A1 20041028	US 2003-743109	20031223			
	A1 20050224	US 2003-743470 US 2003-441795P	20031223			
PRIORITY APPLN. INFO.:			P 20030123			
OTHER SOURCE(S):	MARPAT 141:1905	05				
	lroxy-2,2,12,12-to	etramethyltridecanedi	loic acid			
diethyl ester		((athatia			
RL: PAC (Pharmacolo	gical activity);	RCT (Reactant); SPN	(Synthetic			
preparation); THU	(Therapeutic use)	; BIOL (Biological st	ludy); PREP			
(Preparation); RACT (Reactant or reagent); USES (Uses)						
(anticholesteremic agent; preparation of hydroxyalkanes for cholesterol						
management and related uses)						
RN 738606-33-2 CAPLUS			distant (OGT)			
	i, 7-hydroxy-2,2,	12,12-tetramethyl-, o	netnyl ester (901)			
(CA INDEX NAME)						

738606-34-3P, 7-Hydroxy-2,2,12,12-tetramethyltridecanedioic acid RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(anticholesteremic agent; preparation of hydroxyalkanes for cholesterol management and related uses)

RN 738606-34-3 CAPLUS

CN Tridecanedioic acid, 7-hydroxy-2,2,12,12-tetramethyl- (9CI) (CA INDEX NAME)

GI

Title hydroxyalkanes I [wherein m = 0.5; n = 3.7; X = (CH2)p or CH2; p = 0.5AΒ 0-4; R1, R2, R11, R12 = independently H, alkyl, alkenyl, alkynyl, Ph, PhCH2, wherein R1, R2, R11, and R12 are not simultaneously H; Y1, Y2 = independently alkyl, OH, CO2H, CO2R3, SO3H, (un) substituted heterocyclyl, (di)phosphate and triphosphate esters, etc.; R3 = (un)substituted alkyl, alkenyl, alkynyl, Ph, PhCH2; and pharmaceutically acceptable salts, hydrates, solvates, or mixts. thereof] were prepared for cholesterol management. Thus, reduction of 7-oxo-2,2,12,12-tetramethyltridecanedioic acid di-Et ester with Na(BH4) in MeOH gave 7-hydroxy-2,2,12,12tetramethyltridecanedioic acid di-Et ester (92%), which was saponified with KOH in EtOH to afford the diacid II (95%). The latter inhibited lipid synthesis in primary rat hepatocytes with IC50 of 3.4 μM. In addition, the present invention relates to I, compns. comprising I, and methods useful for treating and preventing a variety of diseases and conditions such as, but not limited to aging, Alzheimer's disease, cancer, cardiovascular disease, diabetic nephropathy, diabetic retinopathy, a disorder of glucose metabolism, dyslipidemia, dyslipoproteinemia,

hypertension, impotence, inflammation, insulin resistance, lipid elimination in bile, obesity, oxysterol elimination in bile, pancreatitis, Parkinson's disease, a peroxisome proliferator activated receptor-associated disorder, phospholipid elimination in bile, renal disease, septicemia, metabolic syndrome disorders (e.g., Syndrome X), thrombotic disorder. Compds. and methods of the invention can also be used to modulate C reactive protein or enhance bile production in a patient (no data). In certain embodiments, I, compns. comprising I, and methods of the invention are useful in combination therapy with other therapeutics, such as hypocholesterolemic and hypoglycemic agents.

L20 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

1987:496416 CAPLUS

DOCUMENT NUMBER:

107:96416

TITLE:

A synthesis of polyoxygenated polycyclic aromatic

compounds via polyketides

AUTHOR (S):

Yamaguchi, Masahiko; Hasebe, Koichi; Shibato, Keisuke;

Nakashima, Hisataka; Minami, Toru

CORPORATE SOURCE:

Dep. Ind. Chem., Kyushu Inst. Technol., Japan

SOURCE:

Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1986),

28th, 627-34 CODEN: TYKYDS

DOCUMENT TYPE:

Journal

LANGUAGE:

Japanese

IT 109873-12-3P 109873-25-8P 109873-26-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT

(Reactant or reagent)

(preparation and intramol. cyclization of)

RN 109873-12-3 CAPLUS

CN Tridecanedioic acid, 7-hydroxy-3,5,9,11-tetraoxo-, dimethyl ester (9CI)

(CA INDEX NAME)

RN 109873-25-8 CAPLUS

CN Tridecanedioic acid, 7-ethyl-7-hydroxy-3,5,9,11-tetraoxo-, dimethyl ester (9CI) (CA INDEX NAME)

RN 109873-26-9 CAPLUS

CN Tridecanedioic acid, 7-hydroxy-3,5,9,11-tetraoxo-7-(3-phenylpropyl)-,

dimethyl ester (9CI) (CA INDEX NAME)

GI

MeO OMe O OH
$$CO_2Me$$
 CO_2Me CO_2Me CH_2CO_2Me III CH_2CO_2Me III CH_2CO_2Me III CH_2CO_2Me CH_2CO_2Me CH_2CO_2Me CH_2CO_2Me CH_2CO_2Me CO_2Me CO_2Me

Ab biomimetic synthesis of polyoxygenated polycyclic aromatic compds. via polyketides was examined Several polyoxoalkanedioates were generated from dicarboxylic acid derivs. and Me acetoacetate dianion. The intramol. condensation of the resulting polyketo esters gave phenolic compds., e.g., indanone I. Ca(OAc)2 was one of the best catalysts. Treatment of aromatic glutarates with the dianion followed by Ca(OAc)2 afforded polyhydroxy derivs. of anthracene and naphthacene related to the natural products. The arenes were oxidized to the corresponding quinones with O2 under basic conditions. The quinones also are aromatic glutarates, and were subjected to further extension of the ring system. Thus, benzanthracene II, prepared from naphthalene III (R = CO2Me, R1 = CH2CO2Me) via polyketide III [R = (COCH2)2CO2Me, R1 = CH2(COCH2)2CO2Me], as outlined above, was converted to dibenzanthracene IV.

L20 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1941:22768 CAPLUS

DOCUMENT NUMBER: 35:22768
ORIGINAL REFERENCE NO.: 35:3608a-g

TITLE: Wound hormones of plants. V. The synthesis of some

analogs of traumatic acid

AUTHOR(S): English, James, Jr.

SOURCE: Journal of the American Chemical Society (1941), 63,

941-3

CODEN: JACSAT; ISSN: 0002-7863

DOCUMENT TYPE: Journal LANGUAGE: Unavailable

IT 855245-29-3, Brassylic acid, ζ-hydroxy-

(preparation of)

RN 855245-29-3 CAPLUS

CN Brassylic acid, ζ-hydroxy- (4CI) (CA INDEX NAME)

AB cf. C. A. 34, 1052.9. The following unsatd. dibasic acids were prepared to test their activity as plant wound hormones. Et sebacate chloride, from the acid ester and SOCl2 in 82% yield, b1 129-30°. The aldehyde esters were prepared from the chlorides by Rosemund's method. Δ1-Unsatd. acids were prepared by adding the desired ester to an equivalent amount of CH2(CO2H)2 in 2-3 times its weight of C5H5N, allowing the mixture to stand 3-5 days, heating on the steam bath for 3 h., acidifying and extracting with ether; the crude mixture of acid esters is hydrolyzed with

N EtOH-NaOH, the acids precipitated with dilute HCl and purified by passing the ether solution through a column packed with an intimate mixture of C and supercel, with final crystallization from dilute EtOH or AcOH; the yields were 30-50%. The $\Delta 2$ -unsatd. acids were prepared as above but with the use of PhNMe2 in place of C5H5N, with sufficient dry MeOH to give a homogeneous reaction mixture; the A2-acids were more soluble in ether and less strongly adsorbed on C than the corresponding $\Delta 1$ -acids. I was also prepared by the method of Bergmann (C. A. 35, 79.7) by using pure azelaic semialdehyde. The following dicarboxylic acids were prepared by these methods: Δ1-nonene-1,9- (I), m. 103°; Δ2-isomer, m. 90°; Δ1-decene-1,10-, m. 165°; Δ2-isomer, m. 109°; Δ1-tridecene-1,13-, m. 108.5°; Δ2-isomer, m. 104°. Heating 40 g. Et α, α' -dibromosebacate and 30 g. PhNMe2 at 180° for 16 h. gives a crude yield of 10 g. of 1,7-octadiene-1,8-dicarboxylic acid, m. 236-9° (decomposition); catalytic reduction of 2 g. with PtO2 (addition of 1 mol. of H) gives 0.6 g. of 1-octene-1,8-dicarboxylic acid, m. 173°. Alkylation of CO(CH2CO2H)2 with PrCHICO2Et according to von Pechman and Sidgwick (Ber. 37, 3816(1904)) (refluxing about 8 h. for each stage), hydrolysis of the material boiling above 150° at 1 mm. by refluxing with 4 times its weight of concentrated HCl for 6-8 h. and crystallization from H2O or dilute EtOH give 52% of

6-undecanone-1,11-dicarboxylic acid (II), m. 114°;
5-nonanone-1,9-dicarboxylic acid, prepared with EtCHICO2H, m. 111°.
Reduction of II with PtO2 and a pressure of 30-40 lb. H in equal vols. of ether and alc. for 2-6 days gives a nearly quant. yield of
6-undecanol-1,11-dicarboxylic acid (III), m. 102-3°;
5-nonanol-1,9-dicarboxylic acid, m. 95°. Heating III with a slight excess of PI3 on the steam bath for 3 h. and refluxing the crude iodo acid with 25% EtOH-KOH for 3 h. give 5-undecene-1,11-dicarboxylic acid, m.
72°; pure 4-nonene-1,9-dicarboxylic acid could not be prepared by this method. All these acids show wound-hormone activity when tested in the presence of the standard co-factor mixture; in all cases the unsatd. acids are more active than the corresponding saturated compds.

=> file reg SINCE FILE TOTAL COST IN U.S. DOLLARS ENTRY SESSION 16.17 871.53 FULL ESTIMATED COST DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION -2.19 -8.03 CA SUBSCRIBER PRICE

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STRUCTURE FILE UPDATES: 25 SEP 2005 HIGHEST RN 863878-84-6 DICTIONARY FILE UPDATES: 25 SEP 2005 HIGHEST RN 863878-84-6

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TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2005

OH

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Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

Uploading C:\Program Files\Stnexp\Queries\107432975.str

chain nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 17 18 19 20 21 23 24
chain bonds :
1-2 1-18 1-19 2-3 3-4 4-5 5-6 6-7 7-8 7-20 8-9 9-10 10-11 11-12 12-13
12-23 12-24 13-14 13-15 15-17 19-21
exact/norm bonds :
1-18 1-19 7-20 13-14 13-15 15-17 19-21
exact bonds :
1-2 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13 12-23 12-24

20

G1:H,Ak

Match level :

1:CLASS 2:CLASS 3:CLASS 4:CLASS 5:CLASS 6:CLASS 7:CLASS 8:CLASS 9:CLASS

10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 17:CLASS 18:CLASS

19:CLASS 20:CLASS 21:CLASS 23:CLASS 24:CLASS

L21 STRUCTURE UPLOADED

=> s 121

SAMPLE SEARCH INITIATED 17:24:41 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 776 TO ITERATE

100.0% PROCESSED 776 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

13849 TO 17191

PROJECTED ANSWERS:

0 TO 0

1.22

0 SEA SSS SAM L21

=> s 121 ful

FULL SEARCH INITIATED 17:24:51 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 15418 TO ITERATE

100.0% PROCESSED 15418 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

L23 2 SEA SSS FUL L21

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 161.33 1032.86

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE 0.00 -8.03

FILE 'CAPLUS' ENTERED AT 17:24:57 ON 26 SEP 2005

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FILE COVERS 1907 - 26 Sep 2005 VOL 143 ISS 14 FILE LAST UPDATED: 25 Sep 2005 (20050925/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 123

L24 1 L23

=> d 124 ibib hitstr abs

L24 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

2004:648491 CAPLUS

DOCUMENT NUMBER:

141:190505

TITLE:

Preparation of hydroxyl compounds for cholesterol

management and related uses

INVENTOR(S):

Dasseux, Jean-Louis Henri; Oniciu, Daniela Carmen

Esperion Therapeutics, Inc., USA

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 348 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE		
WO 2004067489	A2 20040812	WO 2003-US41411	· ·		
WO 2004067489	A3 20041125				
WO 2004067489	C1 20050217				
W: AE, AG, AL,	AM, AT, AU, AZ, B	A, BB, BG, BR, BY,	BZ, CA, CH, CN,		
CO, CR, CU,	CZ, DE, DK, DM, D	Z, EC, EE, EG, ES,	FI, GB, GD, GE,		
GH, GM, HR,	HU, ID, IL, IN, I	S, JP, KE, KG, KP,	KR, KZ, LC, LK,		
LR, LS, LT,	LU, LV, MA, MD, M	G, MK, MN, MW, MX,	MZ, NI, NO, NZ,		
OM, PG, PH,	PL, PT, RO, RU, S	C, SD, SE, SG, SK,	SL, SY, TJ, TM,		
TN, TR, TT,	TZ, UA, UG, US, U	Z, VC, VN, YU, ZA,	ZM, ZW		
RW: BW, GH, GM,	KE, LS, MW, MZ, S	D, SL, SZ, TZ, UG,	ZM, ZW, AM, AZ,		
BY, KG, KZ,	MD, RU, TJ, TM, A	T, BE, BG, CH, CY,	CZ, DE, DK, EE,		
ES, FI, FR,	GB, GR, HU, IE, I	T, LU, MC, NL, PT,	RO, SE, SI, SK,		
TR, BF, BJ,	CF, CG, CI, CM, G	A, GN, GQ, GW, ML,	MR, NE, SN, TD, TG		
US 2004209847	A1 20041021	US 2003-743287	20031223		
US 2004214887	A1 20041028	US 2003-743109	20031223		
US 2005043278	A1 20050224	US 2003-743470	20031223		
PRIORITY APPLN. INFO.:		US 2003-441795P	P 20030123		
OTHER SOURCE(S):	MARPAT 141:190505		! - ! ! -		
IT 738606-33-2P, 7-Hyd	lroxy-2,2,12,12-tet	ramethyltridecaned	ioic acid		
diethyl ester		om (Baratant) ODN	(Comphatia)		
RL: PAC (Pharmacolo	egical activity); R	CT (Reactant); SPN	(Synthetic		
preparation); THU (Therapeutic use);	BIOL (Blological S	cudy); PREP		
(Preparation); RACT (Reactant or reagent); USES (Uses) (anticholesteremic agent; preparation of hydroxyalkanes for cholesterol					
(anticholesteren	ic agent; preparat	tion of hydroxyalka	mes for choresceror		
management and related uses)					
RN 738606-33-2 CAPLUS	; ;	12-totromethyl-	diethyl ester (9CT)		
CN Tridecanedioic acid, 7-hydroxy-2,2,12,12-tetramethyl-, diethyl ester (9CI)					
(CA INDEX NAME)					

738606-34-3P, 7-Hydroxy-2,2,12,12-tetramethyltridecanedioic acid RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(anticholesteremic agent; preparation of hydroxyalkanes for cholesterol management and related uses)

RN 738606-34-3 CAPLUS

CN Tridecanedioic acid, 7-hydroxy-2,2,12,12-tetramethyl- (9CI) (CA INDEX NAME)

GI ·

Title hydroxyalkanes I [wherein m = 0-5; n = 3-7; X = (CH2)p or CH2; p = 0.00AB 0-4; R1, R2, R11, R12 = independently H, alkyl, alkenyl, alkynyl, Ph, PhCH2, wherein R1, R2, R11, and R12 are not simultaneously H; Y1, Y2 = independently alkyl, OH, CO2H, CO2R3, SO3H, (un) substituted heterocyclyl, (di)phosphate and triphosphate esters, etc.; R3 = (un)substituted alkyl, alkenyl, alkynyl, Ph, PhCH2; and pharmaceutically acceptable salts, hydrates, solvates, or mixts. thereof] were prepared for cholesterol management. Thus, reduction of 7-oxo-2,2,12,12-tetramethyltridecanedioic acid di-Et ester with Na(BH4) in MeOH gave 7-hydroxy-2,2,12,12tetramethyltridecanedioic acid di-Et ester (92%), which was saponified with KOH in EtOH to afford the diacid II (95%). The latter inhibited lipid synthesis in primary rat hepatocytes with IC50 of 3.4 μM. In addition, the present invention relates to I, compns. comprising I, and methods useful for treating and preventing a variety of diseases and conditions such as, but not limited to aging, Alzheimer's disease, cancer, cardiovascular disease, diabetic nephropathy, diabetic retinopathy, a disorder of glucose metabolism, dyslipidemia, dyslipoproteinemia,

hypertension, impotence, inflammation, insulin resistance, lipid elimination in bile, obesity, oxysterol elimination in bile, pancreatitis, Parkinson's disease, a peroxisome proliferator activated receptor-associated disorder, phospholipid elimination in bile, renal disease, septicemia, metabolic syndrome disorders (e.g., Syndrome X), thrombotic disorder. Compds. and methods of the invention can also be used to modulate C reactive protein or enhance bile production in a patient (no data). In certain embodiments, I, compns. comprising I, and methods of the invention are useful in combination therapy with other therapeutics, such as hypocholesterolemic and hypoglycemic agents.

=> log y COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	5.39	1038.25
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-0.73	-8.76

STN INTERNATIONAL LOGOFF AT 17:25:48 ON 26 SEP 2005